

#3

OIPE

P. 5

RAW SEQUENCE LISTING
PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001
TIME: 10:09:38

Input Set : A:\ISPH622_Sequence_Listing.txt
Output Set: N:\CRF3\12142001\I005344.raw

ENTERED

4 <110> APPLICANT: Loren J. Miraglia
5 Pamela Nero
6 Mark J. Graham
7 Brett P. Monia
8 Erich Koller
9 MingYi Chiang
10 Mano Manoharan
12 <120> TITLE OF INVENTION: Antisense Modulation of mdm2 expression.
14 <130> FILE REFERENCE: ISPH-0622
C--> 16 <140> CURRENT APPLICATION NUMBER: US/10/005,344
C--> 17 <141> CURRENT FILING DATE: 2001-12-04
19 <150> PRIOR APPLICATION NUMBER: US 09/048,810
20 <151> PRIOR FILING DATE: 1998-03-26
22 <150> PRIOR APPLICATION NUMBER: US 09/280,805
23 <151> PRIOR FILING DATE: 1999-03-26
25 <160> NUMBER OF SEQ ID NOS: 379
27 <170> SOFTWARE: FastSEQ for Windows Version 4.0
29 <210> SEQ ID NO: 1
30 <211> LENGTH: 2372
31 <212> TYPE: DNA
32 <213> ORGANISM: Homo sapiens
34 <220> FEATURE:
35 <221> NAME/KEY: CDS
36 <222> LOCATION: (312)...(1787)
38 <400> SEQUENCE: 1
39 gcaccgcgca agcttggctg cttctggggc ctgtgtggcc ctgtgtgtcg gaaagatgga 60
40 gcaagaagcc gagcccagg ggcggccgca acccctctga ccgagatcct gtcgtttcg 120
41 cagccaggag cacccgtccct ccccgattta gtgcgtacga ggcggccactg ccctggcccc 180
42 gagagtggaa tgcatccccga ggcggccaggc gtcgtgttcc cgcagtagtc agtccccgtg 240
43 aaggaaaactg gggagtcttg agggaccccc gactccaagc gcgaaaaaccc cggatggtga 300
44 ggagcaggca a atg tgc aat acc aac atg tct gta cct act gat ggt gct 350
45 Met Cys Asn Thr Asn Met Ser Val Pro Thr Asp Gly Ala
46 1 5 10
48 gta acc acc tca cag att cca gct tcg gaa caa gag acc ctg gtt aga 398
49 Val Thr Thr Ser Gln Ile Pro Ala Ser Glu Gln Glu Thr Leu Val Arg
50 15 20 25
52 cca aag cca ttg ctt ttg aag tta tta aag tct gtt ggt gca caa aaa 446
53 Pro Lys Pro Leu Leu Lys Leu Leu Lys Ser Val Gly Ala Gln Lys
54 30 35 40 45
56 gac act tat act atg aaa gag gtt ctt ttt tat ctt ggc cag tat att 494
57 Asp Thr Tyr Thr Met Lys Glu Val Leu Phe Tyr Leu Gly Gln Tyr Ile
58 50 55 60
60 atg act aaa cga tta tat gat gag aag caa caa cat att gta tat tgt 542
61 Met Thr Lys Arg Leu Tyr Asp Glu Lys Gln Gln His Ile Val Tyr Cys
62 65 70 75
64 tca aat gat ctt cta gga gat ttg ttt ggc gtg cca agc ttc tct gtg 590
65 Ser Asn Asp Leu Leu Gly Asp Leu Phe Gly Val Pro Ser Phe Ser Val

RAW SEQUENCE LISTING
PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001
TIME: 10:09:38

Input Set : A:\ISPH622_Sequence Listing.txt
Output Set: N:\CRF3\12142001\I005344.raw

66	80	85	90	
68	aaa gag cac agg aaa ata tat acc atg atc tac agg aac ttg gta gta			638
69	Lys Glu His Arg Lys Ile Tyr Thr Met Ile Tyr Arg Asn Leu Val Val			
70	95	100	105	
72	gtc aat cag cag gaa tca tcg gac tca ggt aca tct gtg agt gag aac			686
73	Val Asn Gln Gln Glu Ser Ser Asp Ser Gly Thr Ser Val Ser Glu Asn			
74	110	115	120	125
76	agg tgt cac ctt gaa ggt ggg agt gat caa aag gac ctt gta caa gag			734
77	Arg Cys His Leu Glu Gly Ser Asp Gln Lys Asp Leu Val Gln Glu			
78	130	135	140	
80	ctt cag gaa gag aaa cct tca tct tca cat ttg gtt tct aga cca tct			782
81	Leu Gln Glu Glu Lys Pro Ser Ser His Leu Val Ser Arg Pro Ser			
82	145	150	155	
84	acc tca tct aga agg aga gca att agt gag aca gaa gaa aat tca gat			830
85	Thr Ser Ser Arg Arg Arg Ala Ile Ser Glu Thr Glu Glu Asn Ser Asp			
86	160	165	170	
88	gaa tta tct ggt gaa cga caa aga aaa cgc cac aaa tct gat agt att			878
89	Glu Leu Ser Gly Glu Arg Gln Arg Lys Arg His Lys Ser Asp Ser Ile			
90	175	180	185	
92	tcc ctt tcc ttt gat gaa agc ctg gct ctg tgt gta ata agg gag ata			926
93	Ser Leu Ser Phe Asp Glu Ser Leu Ala Leu Cys Val Ile Arg Glu Ile			
94	190	195	200	205
96	tgt tgt gaa aga agc agt agc agt gaa tct aca ggg acg cca tcg aat			974
97	Cys Cys Glu Arg Ser Ser Ser Glu Ser Thr Gly Thr Pro Ser Asn			
98	210	215	220	
100	ccg gat ctt gat gct ggt gta agt gaa cat tca ggt gat tgg ttg gat			1022
101	Pro Asp Leu Asp Ala Gly Val Ser Glu His Ser Gly Asp Trp Leu Asp			
102	225	230	235	
104	cag gat tca gtt tca gat cag ttt agt gta gaa ttt gaa gtt gaa tct			1070
105	Gln Asp Ser Val Ser Asp Gln Phe Ser Val Glu Phe Glu Val Glu Ser			
106	240	245	250	
108	ctc gac tca gaa gat tat agc ctt agt gaa gaa gga caa gaa ctc tca			1118
109	Leu Asp Ser Glu Asp Tyr Ser Leu Ser Glu Glu Gly Gln Glu Leu Ser			
110	255	260	265	
112	gat gaa gat gat gag gta tat caa gtt act gtg tat cag gca ggg gag			1166
113	Asp Glu Asp Asp Glu Val Tyr Gln Val Thr Val Tyr Gln Ala Gly Glu			
114	270	275	280	285
116	agt gat aca gat tca ttt gaa gaa gat cct gaa att tcc tta gct gac			1214
117	Ser Asp Thr Asp Ser Phe Glu Glu Asp Pro Glu Ile Ser Leu Ala Asp			
118	290	295	300	
120	tat tgg aaa tgc act tca tgc aat gaa atg aat ccc ccc ctt cca tca			1262
121	Tyr Trp Lys Cys Thr Ser Cys Asn Glu Met Asn Pro Pro Leu Pro Ser			
122	305	310	315	
124	cat tgc aac aga tgt tgg gcc ctt cgt gag aat tgg ctt cct gaa gat			1310
125	His Cys Asn Arg Cys Trp Ala Leu Arg Glu Asn Trp Leu Pro Glu Asp			
126	320	325	330	
128	aaa ggg aaa gat aaa ggg gaa atc tct gag aaa gcc aaa ctg gaa aac			1358
129	Lys Gly Lys Asp Lys Gly Glu Ile Ser Glu Lys Ala Lys Leu Glu Asn			
130	335	340	345	

RAW SEQUENCE LISTING
PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001
TIME: 10:09:38

Input Set : A:\ISPH622_Sequence Listing.txt
Output Set: N:\CRF3\12142001\I005344.raw

132 tca aca caa gct gaa gag ggc ttt gat gtt cct gat tgt aaa aaa act	1406
133 Ser Thr Gln Ala Glu Glu Gly Phe Asp Val Pro Asp Cys Lys Lys Thr	
134 350 355 360 365	
136 ata gtg aat gat tcc aga gag tca tgt gtt gag gaa aat gat gat aaa	1454
137 Ile Val Asn Asp Ser Arg Glu Ser Cys Val Glu Asn Asp Asp Lys	
138 370 375 380	
140 att aca caa gct tca caa tca caa gaa agt gaa gac tat tct cag cca	1502
141 Ile Thr Gln Ala Ser Gln Ser Glu Ser Glu Asp Tyr Ser Gln Pro	
142 385 390 395	
144 tca act tct agt agc att att tat agc agc caa gaa gat gtg aaa gag	1550
145 Ser Thr Ser Ser Ile Ile Tyr Ser Ser Gln Glu Asp Val Lys Glu	
146 400 405 410	
148 ttt gaa agg gaa gaa acc caa gac aaa gaa gag agt gtg gaa tct agt	1598
149 Phe Glu Arg Glu Glu Thr Gln Asp Lys Glu Glu Ser Val Glu Ser Ser	
150 415 420 425	
152 ttg ccc ctt aat gcc att gaa cct tgt gtg att tgt caa ggt cga cct	1646
153 Leu Pro Leu Asn Ala Ile Glu Pro Cys Val Ile Cys Gln Gly Arg Pro	
154 430 435 440 445	
156 aaa aat ggt tgc att gtc cat ggc aaa aca gga cat ctt atg gcc tgc	1694
157 Lys Asn Gly Cys Ile Val His Gly Lys Thr Gly His Leu Met Ala Cys	
158 450 455 460	
160 ttt aca tgt gca aag aag cta aag aaa agg aat aag ccc tgc cca gta	1742
161 Phe Thr Cys Ala Lys Lys Leu Lys Lys Arg Asn Lys Pro Cys Pro Val	
162 465 470 475	
164 tgt aga caa cca att caa atg att gtg cta act tat ttc ccc tag	1787
165 Cys Arg Gln Pro Ile Gln Met Ile Val Leu Thr Tyr Phe Pro *	
166 480 485 490	
168 ttgacctgtc tataagagaa ttatatatattt ctaactatat aacccttagga atttagacaa	1847
169 cctgaaattt attcacat atcaaagtga gaaaatgcct caattcacat agatttcttc	1907
170 tcttttagtat aattgaccta ctttggttgt ggaatagtga atacttacta taatttgact	1967
171 tgaatatgtat gctcatcctt tacaccaact cctaattttta aataatttct actctgtctt	2027
172 aaatgagaag tacttggttt tttttttctt aaatatgtat atgacatatta aatgttaactt	2087
173 attatttttt ttgagaccga gtcttgctct gttacccagg ctggagtgca gtgggtgatc	2147
174 ttggctcaact gcaagctctg ccctccccgg gttcgacca ttctcctgcc tcagcctccc	2207
175 aatttagcttgcctacagtc atctgccacc acacctggct aatttttgt acttttagta	2267
176 gagacagggt ttcaccgtgt tagccaggat ggtctcgatc tcctgaccctc gtgatccgcc	2327
177 cacctcgcc tcccaaagtg ctgggattac aggcattgagc caccg	2372
179 <210> SEQ ID NO: 2	
180 <211> LENGTH: 500	
181 <212> TYPE: DNA	
182 <213> ORGANISM: Homo sapiens	
184 <220> FEATURE:	
185 <221> NAME/KEY: misc_signal	
186 <222> LOCATION: (138)...(157)	
187 <223> OTHER INFORMATION: p53 response element RE1	
189 <221> NAME/KEY: misc_signal	
190 <222> LOCATION: (176)...(195)	
191 <223> OTHER INFORMATION: p53 response element RE2	
193 <221> NAME/KEY: exon	

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001

TIME: 10:09:38

Input Set : A:\ISPH622_Sequence Listing.txt
 Output Set: N:\CRF3\12142001\I005344.raw

```

194 <222> LOCATION: (231)...(301)
195 <223> OTHER INFORMATION: Exon 2
197 <221> NAME/KEY: intron
198 <222> LOCATION: (302)...(422)
199 <223> OTHER INFORMATION: Intron 2
201 <400> SEQUENCE: 2
202 ggctgcgggc ccctgcggcg cgggagggtcc ggatgatcgc aggtgcctgt cgggtcaacta 60
203 gtgtgaacgc tgcgcgtagt ctggggcggga ttggggccgt tcagtgccaa ggttgcactca 120
204 gcttttcctc tttagctgggt caagttcaga cacgttccga aactgcagta aaaggagtttta 180
205 agtcctgact tgtctccagc tggggctatt taaaccatgc attttccag ctgtgttcag 240
206 tggcgattgg agggttagacc tggggccacg gacgcacgccc accttttctc tgctgatcca 300
207 gttaagcacc gacttgcttg tagcttttagt tttaactgtt gtttatgtt tttatataatg 360
208 atgtattttc cacagatgtt tcatgatttc cagttttcat cgtgtctttt tttcccttgt 420
209 aggcaaatgt gcaataccaa catgtctgtt cctactgtt gggctgttaac caccacacag 480
210 attccagctt cggaacaaga 500
212 <210> SEQ ID NO: 3
213 <211> LENGTH: 20
214 <212> TYPE: DNA
215 <213> ORGANISM: Artificial Sequence
217 <220> FEATURE:
218 <223> OTHER INFORMATION: Antisense Oligonucleotide
220 <400> SEQUENCE: 3
221 cagccaagct cgcgcgggtc 20
223 <210> SEQ ID NO: 4
224 <211> LENGTH: 20
225 <212> TYPE: DNA
226 <213> ORGANISM: Artificial Sequence
228 <220> FEATURE:
229 <223> OTHER INFORMATION: Antisense Oligonucleotide
231 <400> SEQUENCE: 4
232 tctttccgac acacaggggcc 20
234 <210> SEQ ID NO: 5
235 <211> LENGTH: 20
236 <212> TYPE: DNA
237 <213> ORGANISM: Artificial Sequence
239 <220> FEATURE:
240 <223> OTHER INFORMATION: Antisense Oligonucleotide
242 <400> SEQUENCE: 5
243 cagcaggatc tcggtcagag 20
245 <210> SEQ ID NO: 6
246 <211> LENGTH: 20
247 <212> TYPE: DNA
248 <213> ORGANISM: Artificial Sequence
250 <220> FEATURE:
251 <223> OTHER INFORMATION: Antisense Oligonucleotide
253 <400> SEQUENCE: 6
254 gggcgctcgat acgcactaat 20
256 <210> SEQ ID NO: 7
257 <211> LENGTH: 20

```

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001

TIME: 10:09:38

Input Set : A:\ISPH622_Sequence Listing.txt
 Output Set: N:\CRF3\12142001\I005344.raw

```

258 <212> TYPE: DNA
259 <213> ORGANISM: Artificial Sequence
261 <220> FEATURE:
262 <223> OTHER INFORMATION: Antisense Oligonucleotide
264 <400> SEQUENCE: 7
265 tcggggatca ttccactctc 20
267 <210> SEQ ID NO: 8
268 <211> LENGTH: 20
269 <212> TYPE: DNA
270 <213> ORGANISM: Artificial Sequence
272 <220> FEATURE:
273 <223> OTHER INFORMATION: Antisense Oligonucleotide
275 <400> SEQUENCE: 8
276 cgggggtttc gcgcttggag 20
278 <210> SEQ ID NO: 9
279 <211> LENGTH: 20
280 <212> TYPE: DNA
281 <213> ORGANISM: Artificial Sequence
283 <220> FEATURE:
284 <223> OTHER INFORMATION: Antisense Oligonucleotide
286 <400> SEQUENCE: 9
287 catttgccctg ctcctcacca 20
289 <210> SEQ ID NO: 10
290 <211> LENGTH: 20
291 <212> TYPE: DNA
292 <213> ORGANISM: Artificial Sequence
294 <220> FEATURE:
295 <223> OTHER INFORMATION: Antisense Oligonucleotide
297 <400> SEQUENCE: 10
298 gtattgcaca tttgcgtgtct 20
300 <210> SEQ ID NO: 11
301 <211> LENGTH: 20
302 <212> TYPE: DNA
303 <213> ORGANISM: Artificial Sequence
305 <220> FEATURE:
306 <223> OTHER INFORMATION: Antisense Oligonucleotide
308 <400> SEQUENCE: 11
309 agcaccatca gtaggtacag 20
311 <210> SEQ ID NO: 12
312 <211> LENGTH: 20
313 <212> TYPE: DNA
314 <213> ORGANISM: Artificial Sequence
316 <220> FEATURE:
317 <223> OTHER INFORMATION: Antisense Oligonucleotide
319 <400> SEQUENCE: 12
320 ctaccaaggc cctgttagatc 20
322 <210> SEQ ID NO: 13
323 <211> LENGTH: 20
324 <212> TYPE: DNA

```

Use of n and / or Xaa has been detected in the Sequence Listing. Review the Sequence Listing to ensure a corresponding explanation is present in the <220> to <223> fields of each sequence using n or Xaa.

VERIFICATION SUMMARY

PATENT APPLICATION: US/10/005,344

DATE: 12/14/2001

TIME: 10:09:39

Input Set : A:\ISPH622_Sequence Listing.txt

Output Set: N:\CRF3\12142001\I005344.raw

L:16 M:270 C: Current Application Number differs, Replaced Application Number

L:17 M:271 C: Current Filing Date differs, Replaced Current Filing Date

L:4424 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:370

L:4425 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:370

L:4536 M:334 W: (2) Invalid Amino Acid in Coding Region, NUMBER OF INVALID KEYS:3